What is claimed is:

1. A compound of the following formula I, or an enantiomer, diastereomer, salt or metabolite thereof:

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wherein:

 $R_{2} \ is \ hydrogen, \ halogen, \ -CHO, \ alkyl, \ haloalkyl, \ (cycloalkyl)alkyl, \ alkenyl, \\ alkynyl, \ alkoxyalkyl, \ haloalkoxyalkyl, \ alkoxy, \ aryloxy \ alkoxyalkoxy, \\ cyano, \ hydroxy, \ hydroxyalkyl, \ nitro, \ -CH(OR_{_{13}})(OR_{_{14}}), \ -(CH_{_2})_{_{w}}Y; \\ with \ the \ proviso \ that \ when \ R_{_1} \ is \ \underline{B}, \ R_{_2} \ is \ not \ hydrogen, \ halogen, \\ alkyl, \ haloalkyl, \ alkoxy, \ hydroxyalkyl, \ nitro, \ -(CH_{_2})_{_{w}}NR_{_{19}}R_{_{20}} \ or \\ -NHSO_{_2}R_{_{22}};$

10 R₃ is heteroaryl;

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 R_4 and R_5 are each independently alkyl, hydroxyalkyl, cycloalkyl, hydroxy substituted cycloalkyl, alkoxyalkyl, or hydroxy substituted alkoxyalkyl, or R_4 and R_5 together form a cyclobutyl, cyclopentyl, cyclohexyl, tetrahydrofuranyl or tetrahydropyranyl ring which may be optionally substituted with one or more hydroxy groups;

 R_6 is alkyl, hydroxyalkyl, haloalkyl, hydroxy substituted haloalkyl, cycloalkyl, hydroxy substituted cycloalkyl, (cycloalkyl)alkyl, hydroxy substituted (cycloalkyl)alkyl, aralkyl, alkoxy, hydroxy substituted alkoxy, alkoxyalkyl, hydroxy substituted alkoxyalkyl, or $-NR_{16}R_{17}$;

 $R_{7} \text{ is } -(CH_{2})_{w} - CO_{2}R_{15}, -(CH_{2})_{w} - (C=O)NR_{16}R_{17}, -(CH_{2})_{w} - NR_{15}(C=O)NR_{16}R_{17}, - \\ (CH_{2})_{w} - CH_{2}OH, -(CH_{2})_{w} - (C=O)R_{15}, \text{ tetrazolyl, oxadiazolyl or triazolyl } \\ \text{wherein said tetrazolyl, oxadiazolyl or triazolyl may optionally be} \\ \text{substituted with hydrogen, alkyl, hydroxy or halogen;}$

- R_{s} , R_{g} , R_{ga} , R_{10} and R_{12} are each independently hydrogen, halogen, alkyl, hydroxyalkyl, cycloalkyl, (cycloalkyl)alkyl, aryl, heteroaryl, arylalkyl, alkylthioalkyl, alkoxy or alkoxyalkyl, or R_{g} and R_{ga} together with the carbon atom to which they are bonded form a cycloalkyl ring;
- 10 $R_{_{11}}$ and $R_{_{11a}}$ are each independently hydrogen, alkoxy, or together form a carbonyl;

 $R_{{\scriptscriptstyle 13}}$ and $R_{{\scriptscriptstyle 14}}$ are alkyl or together form a five to six-membered ring;

 R_{15} , R_{16} and R_{17} are independently hydrogen, alkyl, hydroxyalkyl, cycloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, aralkyl, heterocycloalkyl, aryl, heteroaryl or -(CH $_2$) $_w$ Q, or R_{16} and R_{17} may together form a four to six–membered heterocyclic ring;

n is 1 or 2;

15

w is 0, 1, or 2;

Y is heteroaryl, -COOH, -COOR₁₈, -CONR₁₉R₂₀, -NR₁₉R₂₀, -NR₁₉-OR₂₀, -20 $NR_{21}(C=O)R_{22}, -NR_{21}(C=O)NR_{19}R_{20}, -N(R_{19})-(alk)-NR_{21}(C=O)R_{22},$ $-NR_{21}(C=O)OR_{18}, -NR_{21}SO_{2}R_{22}, -SO_{2}R_{22}, \underline{Q}, \underline{R} \text{ or } \underline{S};$

$$Q \text{ is }$$
 R_{23}
 R_{24}
 X

$$\frac{R_{23}}{Z} \stackrel{R_{24}}{\nearrow}$$

 $R_{_{18}}$, $R_{_{19}}$, $R_{_{20}}$, $R_{_{21}}$ and $R_{_{22}}$ are each independently hydrogen, alkyl, haloalkyl, alkoxyalkyl, cycloalkyl, alkenyl, alkynyl, aryl, aralkyl, heteroaryl, or $R_{_{19}}$ and $R_{_{20}}$ may together form a four to seven-membered heterocyclic ring;

 R_{23} and R_{24} are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;

Z is oxygen,
$$N-R_{25}$$
 or R_{26}

x is 2, 3 or 4;

5

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10 R_{25} , R_{26} and R_{27} are each independently hydrogen, alkyl or cycloalkyl, or R_{26} and R_{27} may together form a three to seven-membered cycloalkyl ring;

 R_{101} , R_{102} , R_{103} , and R_{104} are each independently hydrogen, halogen, -CHO, alkyl, haloalkyl, (cycloalkyl)alkyl, alkenyl, alkynyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, cyano, hydroxy, hydroxyalkyl, nitro, -CH(OR₁₃)(OR₁₄), or -(CH₂) $_{\rm w}$ Y;

wherein said rings; aryl alone or as part of another group; or heteroaryl alone or as part of another group may each optionally be substituted by one or more hydrogen, halogen, cyano, alkyl, hydroxyalkyl, alkoxy, nitro or trifluoromethyl groups;

provided that when R_i is \underline{A} said compound is other than

2. A compound of claim 1, wherein

$$R_{1} \text{ is } \underbrace{\underline{A}}_{R_{1}} , \underbrace{\underline{D}}_{R_{1}} , \underbrace{\underline{R}_{8}}_{R_{1}} , \underbrace{\underline{R}_{8}}_{R_{1}} , \underbrace{\underline{R}_{10}}_{R_{10}}$$

<u>M</u>

5

R₂ is alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or $-(CH_2)_wY$, or when R_1 is \underline{D} , R_2 is hydrogen, alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or -(CH₉), Y;

10

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- R₃ is isoxazolyl pyridizinyl, pyrazinyl or pyrimidinyl, each optionally independently substituted with one to three substituents selected from hydrogen, halogen, cyano, alkyl, alkoxy, trifluoromethyl or nitro;
- 15 R_4 and R_5 are each independently alkyl, cycloalkyl, or R_4 and R_5 together form a cyclobutyl, cyclopentyl or cyclohexyl ring;

R₆ is alkyl, haloalkyl, cycloalkyl or alkoxy;

 R_7 is $-CO_2R_{15}$, $-(C=O)NR_{16}R_{17}$ or $-CH_2OH$;

 $R_{\mbox{\tiny 8}},\,R_{\mbox{\tiny 9}},\,R_{\mbox{\tiny 10}}$ and $R_{\mbox{\tiny 12}}$ are each independently hydrogen, halogen, alkyl, cycloalkyl, alkoxy or alkoxyalkyl;

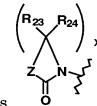
 R_{11} and R_{11a} are each independently hydrogen, alkoxy, or together form a carbonyl;

 $R_{\scriptscriptstyle 15},\,R_{\scriptscriptstyle 16}$ and $R_{\scriptscriptstyle 17}$ are independently hydrogen, alkyl or cycloalkyl or $R_{\scriptscriptstyle 16}$ and R_{17} may together form a four to six-membered heterocyclic ring;

5 n is 1 or 2;

w is 0, 1, or 2;

Y is $-COOR_{18}$, $-NR_{21}(C=O)R_{22}$, $-NR_{21}(C=O)NR_{19}R_{20}$, $-NR_{21}(C=O)OR_{18}$, $-NR_{21}SO_2R_{22}$, $-SO_2R_{22}$ or Q;



10 R_{18} , R_{19} , R_{20} , R_{21} and R_{22} are each independently hydrogen, alkyl, cycloalkyl, or R_{19} and R_{20} may together form a four to seven-membered heterocyclic ring;

 R_{23} and R_{24} are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;

$$N-R_{25}$$
 R_{26}

15 Z is oxygen,

x is 2, 3 or 4;

 R_{25} , R_{26} and R_{27} are each independently hydrogen, alkyl or cycloalkyl, or R_{26} and R_{27} may together form a three to seven-membered cycloalkyl ring;

20 R_{101} , R_{102} , R_{103} , and R_{104} are each independently hydrogen, halogen, alkoxy or alkyl.

> 3. A compound of claim 1, wherein

 R_2 is alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, hydroxyalkyl, or $-(CH_2)_wY$, or when R_1 is \underline{D} , R_2 is hydrogen, alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxyalkoxy, hydroxyalkyl, or $-(CH_2)_wY$;

 $R_{\scriptscriptstyle 3}$ is isoxazolyl, optionally independently substituted with one or two substituents selected from hydrogen, halogen, cyano, alkyl, alkoxy, trifluoromethyl or nitro;

 R_4 and R_5 are each independently alkyl, cycloalkyl, or R_4 and R_5 together form a cyclobutyl, cyclopentyl or cyclohexyl ring;

R₆ is alkyl, haloalkyl, cycloalkyl or alkoxy;

 $R_{_{7}}$ is - $CO_{_{2}}R_{_{15}}$ or -(C=O)N $R_{_{16}}R_{_{17}}$;

15 R_8 , R_9 and R_{10} are each independently hydrogen, halogen, alkyl, cycloalkyl, alkoxy or alkoxyalkyl;

 $\boldsymbol{R}_{_{11}}$ and $\boldsymbol{R}_{_{11a}}$ together form a carbonyl;

 $R_{_{15}}$, $R_{_{16}}$ and $R_{_{17}}$ are independently hydrogen, alkyl, or cycloalkyl or $R_{_{16}}$ and $R_{_{17}}$ may together form a four to six–membered heterocyclic ring;

20 n is 2;

5

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w is 0, 1, or 2;

Y is -NR₂₁(C=O)R₂₂, -NR₂₁(C=O)NR₁₉R₂₀, -NR₂₁(C=O)OR₁₈, -NR₂₁SO₂R₂₂, -SO₂R₂₂ or **Q**;

$$\begin{array}{c|c}
\begin{pmatrix}
R_{23} & R_{24} \\
Z & N
\end{pmatrix}$$

$$Q \text{ is}$$

 R_{18} , R_{19} , R_{20} , R_{21} and R_{22} are each independently hydrogen, alkyl, cycloalkyl, or R_{19} and R_{20} may together form a four to seven-membered heterocyclic ring;

5 R_{23} and R_{24} are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;

$$N-R_{25}$$
 or R_{26}

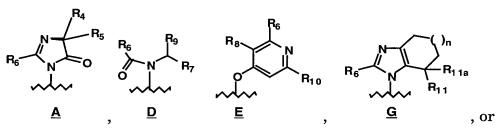
Z is oxygen,

x is 2, 3 or 4;

 R_{25} , R_{26} and R_{27} are each independently hydrogen, alkyl or cycloalkyl, or R_{26} and R_{27} may together form a three to seven-membered cycloalkyl ring;

 $R_{101},\,R_{102},\,R_{103},$ and R_{104} are each independently hydrogen, halogen, or alkyl.

4. A compound of claim 1, wherein



$$\begin{array}{c|c} & & & \\ & & & \\ R_6 & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ &$$

R, is

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 R_2 is alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl,or $-(CH_2)_wY$; or when R_1 is \underline{D} , R_2

is hydrogen, alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or $-(CH_2)_wY$;

 R_3 is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen;

 R_4 and R_5 are each independently alkyl, cycloalkyl, or R_4 and R_5 together form a cyclobutyl, cyclopentyl or cyclohexyl ring;

R₆ is alkyl, haloalkyl, cycloalkyl or alkoxy;

 R_7 is $-(C=O)NR_{16}R_{17}$;

10 R_8 , R_9 , and R_{10} are independently H, alkyl, cycloalkyl, alkoxy or alkoxyalkyl;

n is 2;

w is 0, 1, or 2;

 $Y \ is \ -NR_{_{21}}(C=O)R_{_{22}}, \ -NR_{_{21}}(C=O)NR_{_{19}}R_{_{20}}, \ -NR_{_{21}}(C=O)OR_{_{18}}, \ -NR_{_{21}}SO_{_2}R_{_{22}}$

15 or Q;

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$$Q \text{ is } Q \text{ of } Q \text{$$

 R_{18} , R_{19} , R_{20} , R_{21} and R_{22} are each independently hydrogen, alkyl, cycloalkyl, or R_{19} and R_{20} may together form a four-, five-, six- or to seven-membered heterocyclic ring;

 R_{23} and R_{24} are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;

$$Z \text{ is}$$
 $N-R_{25}$ or R_{26}

x is 2;

 R_{25} , R_{26} and R_{27} are each independently hydrogen, alkyl or cycloalkyl, or R_{26} and R_{27} may together form a three-, four-, five, six- or sevenmembered cycloalkyl ring;

5

 R_{101} , R_{102} , R_{103} , and R_{104} are each independently hydrogen, halogen, or alkyl.

5. A compound of claim 1, wherein R_3 is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.

- 6. A compound of claim 5, wherein R_1 is
- 7. A compound of claim 6, wherein R_2 is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R_{101} , R_{102} , R_{103} , R_{104} are each independently hydrogen, halogen, or alkyl.
 - 8. A compound of claim 6, wherein R_2 is $-CH_2Y$.
- 9. A compound of claim 8, wherein Y is Q.

- 10. A compound of claim 5, wherein R_1 is
- 11. A compound of claim 10, wherein R_2 is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R_{101} , R_{102} , R_{103} , R_{104} are each independently hydrogen, halogen, or alkyl.
 - 12. A compound of claim 10, wherein R₂ is -CH₂Y.
 - 13. A compound of claim 12, wherein Y is Q.

- 14. A compound of claim 5, wherein R_1 is
- 15. A compound of claim 14, wherein R_2 is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R_{101} , R_{102} , R_{103} , R_{104} are each independently hydrogen, halogen, or alkyl.
 - 16. A compound of claim 14, wherein R_2 is $-CH_2Y$.
- 10 17. A compound of claim 16, wherein Y is Q.
 - 18. A compound of claim 1, wherein R_2 is alkoxyalkyl alkyl, haloalkyl or haloalkoxyalkyl
- 15 19. A compound of claim 18, wherein R_3 is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.
 - 20. A compound of claim 1, wherein R_2 is $-CH_2Y$.
- 21. A compound of claim 20, wherein R_3 is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.
- 25 22. A compound of claim 20, wherein Y is Q.

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- 23. A compound of claim 22, wherein $R_{\scriptscriptstyle 3}$ is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.
- 5 24. A compound of claim 1, wherein said compound is selected from the group consisting of
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(methylamino)methyl][1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3-methyl-2-oxo-1-imidazolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
 - $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(2-pyrazinyl)~[1,1'-biphenyl]-2-sulfonamide;$
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3-chloro-2-pyrazinyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(3,6-dimethyl-2-pyrazinyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(3-methoxy-2-pyrazinyl)[1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
 - (S)-N-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-[2-methyl-1-(3-methyl-1,2,4-oxadiazol-5-yl)propyl]pentanamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
 - N-(3,4-Dimethyl-5-isoxazolyl)-4'-[2-(2-methoxyethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
 - N-(3,4-Dimethyl-5-isoxazolyl)-4'-[2-(ethoxymethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'[(2-oxo-1-pyrrolidinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]2-sulfonamide;
- 4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'[(3-methyl-2-oxo-1-imidazolidinyl)methyl]-N-(3,4-dimethyl-5isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
 - (S)-2-[N-[2'-[[N-(3-Methyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;
- 30 (S)-2-[N-[2'-[[N-(4-Bromo-3-methyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;

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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]
                  -N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
                                       4'-[(7-Methoxycarbonyl-2-ethoxybenzimidazol-1-yl)methyl]-
                  N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
                                         4'-[(7-Carboxy-2-ethoxybenzimidazol-1-yl)methyl]-N-
   5
                  (3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
                                         4'-[(7-Methoxycarbonyl-2-ethylbenzimidazol-1-yl)methyl]-N-
                  (3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
                                       4'-[(7-Carboxy-2-ethylbenzimidazol-1-yl)methyl]-N-
10
                  (3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
                                       2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-
                  (methoxycarbonyl)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-
                  isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
                                        2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4'-[(2-ethoxy-7-indin-1-yl)methyl]-4
                  (carboxy) benzimidazol-1-yl) methyl]-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-isoxazolyl)-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazolyl]-[1,1'-isoxazoly
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                  biphenyl]-2-sulfonamide;
                                       2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N-
                  methylcarbamoyl)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-
                   [1,1'-biphenyl]-2-sulfonamide;
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                                        2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N,N-
                  dimethylcarbamoyl)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-
                  isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
                                        4'-[(2-Ethylquinolin-4-yl)oxymethyl]-N-(1,3,5-trimethylpyrazol-4-yl)
                     [1,1'-biphenyl]-2-sulfonamide;
                                        4'-[(2-Ethylquinolin-4-yl)oxymethyl]-N-(3-methylisoxazol-5-yl) [1,1'-
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                  biphenyl]-2-sulfonamide;
                                        4'-[(5-Acetyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(3,4-
                   dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
                                        4'-[(5-Methoxycarbonyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-
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                  (3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
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- 4'-[(5-(N,N-dimethylcarbamoyl)-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-hydroxymethyl[1,1'-biphenyl]-2-sulfonamide
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-10 dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(3-methoxy-5-methyl-2-pyrazinyl)[1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4-bromo-3-methyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(formylmethylamino)-methyl][1,1'-biphenyl]-2-sulfonamide;
 - $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylpropanamide;$
 - N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethylcyclopropanecarbox-amide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-25 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,2-dimethylpropanamide;
 - N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethylbutanamide;

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 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2-methoxy-N-methylacetamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethyl-4-pentynamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\ [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylcyclobutanecarboxamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,3-dimethylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,2,2-trimethylpropanamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3-methoxy-N-methylpropanamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2ethoxy-N-methylacetamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methyl-2-furancarboxamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,4-dimethylpentanamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylbenzamide;$

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 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\ [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methyl-3-thiophenecarboxamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethylcyclopentaneacetamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\ [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylcyclohexanecarboxamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,3-dimethylbenzamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\ [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylbenzeneacetamide;$

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2-fluoro-N-methylbenzamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3fluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-4fluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethylcyclohexaneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2-fluoro-N-methylbenzeneacetamide;

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N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3-
fluoro-N-methylbenzeneacetamide;
      N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-4-
fluoro-N-methylbenzeneacetamide;
      N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-
N,N',N'-trimethylurea;
      N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N'-
(1,1-dimethylethyl)-N-methylurea;
       [[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
yl]methyl]methylcarbamic acid ethyl ester;
       \hbox{\tt [[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-}\\
[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
yl]methyl]methylcarbamic acid 2-methylpropyl ester;
      N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-
N,3,3-trimethylbutanamide;
      N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-
methyl-2-pyridinecarboxamide;
      N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-
methyl-3-pyridinecarboxamide;
      N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
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N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethyl-2-pyrazinecarboxamide;

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 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,1-dimethyl-1H-pyrrole-2-carboxamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethyl-1,2,3-thiadiazole-4-carboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,5-dimethyl-2-pyrazinecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,3,5-trimethyl-4-isoxazolecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,3-dimethyl-2-thiophenecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,5-dimethyl-2-thiophenecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3cyano-N-methylbenzamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\ [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-4-cyano-N-methylbenzamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2methoxy-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2chloro-N-methylbenzamide;

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 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\ [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3-chloro-N-methylbenzamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-4chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2,3-difluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]3,4-difluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]3,5-difluoro-N-methylbenzamide;

4-Acetyl-N-[[4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3ethoxy-N-methyl-2-thiophenecarboxamide;

 $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;$

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(propylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;

N-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(1-oxopentyl)-L-valine methyl ester;

N-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-30 yl]methyl]-N-(1-oxopentyl)-L-valine;

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      N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-
pyrrolidinyl)methyl]-4'-[(2-propyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-
yl)methyl][1,1'-biphenyl]-2-sulfonamide;
      N-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-
dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-
yl]methyl]-N,3,3-trimethylbutanamide;
      N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
4-yl]methyl]-N2-(1-oxopentyl)-L-valinamide;
      N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
      N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
4-yl]methyl]-N,N-dimethyl-N2-(1-oxopentyl)-L-valinamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
dimethyl-5-isoxazolyl)-2'-[[(2,2,2-trifluoroethyl)amino]methyl][1,1'-
biphenyl]-2-sulfonamide;
dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
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- 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(3,4-
- dimethyl-5-isoxazolyl)-2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;
- 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(3,4dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid methyl ester;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4dimethyl-5-isoxazolyl)-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4dimethyl-5-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;
 - 2'-(Cyanomethyl)-N-(3,4-dimethyl-5-isoxazolyl)-4'-[(2-ethyl-5,7dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4,4]non-1-en-3-yl)methyl]-2'-(cyanomethyl)-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-cyano-
     N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
           4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
     dimethyl-5-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
 5
           2'-Cyano-N-(3,4-dimethyl-5-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-
     imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
           N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-methyl[1,1'-
     biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
           N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-
     imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[[(2,2,2-
10
     trifluoroethyl)amino]methyl][1,1'-biphenyl]-2-sulfonamide;
           N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
     [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
     yl]benzeneacetamide;
           15
     [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]-3,3-
     dimethylbutanamide;
           2'-Amino-4'-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-
     N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
           4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
20
     dimethyl-5-isoxazolyl)-2'-nitro[1,1'-biphenyl]-2-sulfonamide;
           N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-isoleucinamide;
           N2-(Cyclopropylcarbonyl)- N2-[[2'-[[(3,4-dimethyl-5-
     isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
25
     isoleucinamide;
           N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-isoleucinamide;
           N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yllmethyll-N-methyl- N2-(3-methyl-1-oxobutyl)-L-isoleucinamide;
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N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-isoleucinamide;
            N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-isoleucinamide;
 5
            N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-leucinamide;
            N2-(Cyclopropylcarbonyl)- N2-[[2'-[[(3,4-dimethyl-5-
     isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
     leucinamide;
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            N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-leucinamide;
            N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(phenylacetyl)-L-leucinamide;
            N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
15
     4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-leucinamide;
            N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-leucinamide;
           N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-leucinamide;
            N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
20
     4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-valinamide;
            N2-(Cyclopropylcarbonyl)- N2-[[2'-[[(3,4-dimethyl-5-
     isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
     valinamide;
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           N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-valinamide;
           N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
     4-yl]methyl]-N-methyl- N2-(phenylacetyl)-L-valinamide;
            N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
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     4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-valinamide;
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N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-valinamide;
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N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-valinamide;

5 N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-ethyl- N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-hexyl- N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-cyano[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-hydroxymethyl[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;

N, N-Dimethyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl) methyl]-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;

N-Methyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

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 $N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-\\ (methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;$

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;

2'-Chloro-N-(3,4-dimethyl-5-isoxazolyl)-4'-[[(5,6,7,8-tetrahydro-2-ethyl-4-quinolinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-fluoro[1,1'-

biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

 $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(phenoxymethyl)[1,1'-biphenyl]-2-sulfonamide;$

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-(Cyclopropylcarbonyl)-N2-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

N2-(Cyclopropylcarbonyl)- N2-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-

25 (methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

N2-[[2-Chloro-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-

30 (trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

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N2-(Cyclobutylcarbonyl)-N2-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-valinamide;
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- 1-[[2-Chloro-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxylic acid;
 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(methylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;
- (S)-N-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-10 biphenyl]-4-yl]methyl]-N-[2-methyl-1-[(4-methyl-1-piperazinyl)carbonyl]propyl]pentanamide;
 - $(S)-N-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(1-piperidinyl)carbonyl]propyl]pentanamide; \\ N-(3,3-Dimethylbutyl)- N2-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methylbutyl)- N2-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methylbutyl)- N2-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyll-5-isoxazolyl)amino[sulfonyl][1,1'-biphenyl]-4-yl]methyll-N-(1-piperidinyl)carbonyl]propyl]pentanamide; \\ N-(3,3-Dimethylbutyl)- N2-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-1-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-1-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-1-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-1-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-1-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-1-[(3,4-dimethyl-5-isoxazolyl)amino[sulfonyl]-$
- isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-[(4-fluorophenyl)methyl]-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(methylethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(propoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4-Chloro-1-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-

biphenyl]-4-yl]methyl]-2-propyl-1H-imidazole-5-carboxamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

 $4\text{'-}[(2\text{-Butyl-4-oxo-1,3-diazaspiro}[4.4]non-1\text{-en-3-yl})methyl]-2\text{'-}[(1,2\text{-dihydro-2-oxo-1-pyridinyl})methyl]-N-(3,4\text{-dimethyl-5-isoxazolyl})[1,1\text{'-dihydro-2-oxo-1-pyridinyl})methyl]-N-(3,4\text{-dimethyl-5-isoxazolyl})[1,1\text{'-dimethyl-5-isoxazolyl})[1,1\text{'-dimethyl-5-isoxazolyl})[1,1\text{'-dimethyl-5-isoxazolyl})[1,1\text{'-dimethyl-5-isoxazolyl})[1,1\text{'-dimethyl-5-isoxazolyl}][1,1\text{'-dimethyl-5-isoxazolyl][1,1\text{'-dimethyl-5-isoxazolyl]}[1,1\text{'-dimethyl-5-isoxazolyl][1,1\text{'-dimethyl-5-isoxazolyl]}[1,1\text{'-dimethyl-5-isox$

30 biphenyl]-2-sulfonamide;

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dimethyl-5-isoxazolyl)-2'-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-2-
                    sulfonamide;
                                          2-Butyl-4-chloro-1-[[2'-[[(3,4-dimethyl-5-
                    isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-1H-imidazole-5-
     5
                    carboxamide;
                                          N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2-methyl-4-isoxazolyl)-4'-[(2-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazolyl)-4'-[(3-methyl-4-isoxazoly
                    quinolinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
                                          N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2-ethyl-4-quinolinyl)oxy]methyl]
 10
                    [1,1'-biphenyl]-2-sulfonamide;
                                          N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2-ethyl-5,6,7,8-tetrahydro-4-
                    quinolinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
                                          N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2-propyl-4-
                    quinolinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
15
                                          N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(5,6,7,8-tetrahydro-2,4-dimethyl-7-
                    oxopyrido[2,3-d]pyrimidin-8-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
                                          4'-[[(2-Ethyl-4-quinolinyl)oxy]methyl]-N-(3,4-dimethyl-5-isoxazolyl)-
                    2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-
                   sulfonamide;
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                                         N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxaz
                   pyrrolidinyl)methyl]-4'-[[(5,6,7,8-tetrahydro-2-ethyl-4-
                   quinolinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
                                          3-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
                   yl]methyl]-2-ethyl-N-methyl-3H-benzimidazole-4-carboxamide;
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                                          1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
                   yl]methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid phenylmethyl ester;
                                          1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
                   yl]methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-phenylethyl ester;
                                         1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
30
                   yl]methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-(2-oxo-1-
                   pyrrolidinyl)ethyl ester;
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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-

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1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 3-(2-oxo-1-
pyrrolidinyl)propyl ester;
      2'-Cyano-N-(3,4-dimethyl-5-isoxazolyl)-4'-[[(2-ethyl-4-
quinolinyl)oxy[methyl][1,1'-biphenyl]-2-sulfonamide;
      2'-(Cyanomethyl)-N-(3,4-dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-
hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-
sulfonamide;
      3-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-5-ethyl-N-methyl-2-propyl-3H-imidazole-4-carboxamide;
      1-[[2-Chloro-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-
biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
      3-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-5-ethyl-2-propyl-3H-imidazole-4-carboxamide;
      3-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-2-ethoxy-N-methyl-3H-benzimidazole-4-carboxamide;
      3-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-2-ethoxy-N,N-dimethyl-3H-benzimidazole-4-carboxamide;
      2-[[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]propylamino]-3-pyridinecarboxylic acid;
      4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(3,4-dimethyl-5-
isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
      N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-
propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
      4'-[(2,7-Diethyl-5H-pyrazolo[1,5-b][1,2,4]triazol-5-yl)methyl]-N-(3,4-
dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
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N-[2-Butyl-3-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-3,4-dihydro-4-oxo-6-quinazolinyl]-N'-methyl-N'-(1-methylethyl)urea;

2-[[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl] methyl] propylamino]-N-methyl-3-pyridinecarboxamide;

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4'-[[(3-Methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

3-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethyl-5-methyl-3H-imidazole-4-carboxamide;

4'-[(2-Butyl-3,4-dihydro-4-oxo-3-quinazolinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(4,4-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-[2-[[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl]methyl]methylamino]ethyl]acetamide;

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4-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-acetic acid ethyl ester;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N2-(1-oxopentyl)-N-propyl-L-valinamide;

N2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N2-(1-oxopentyl)-N-[(tetrahydro-2-furanyl)methyl]-L-valinamide;

2'-Chloro-N-(3,4-dimethyl-5-isoxazolyl)-4'-[[(2-ethyl-4-quinolinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2-ethyl-4-quinolinyl)oxy]methyl]-2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-chloro-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2-methylpropoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(ethylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;

 $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2,2,2-trifluoroethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;$

 $\label{lem:conditional} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2-fluoroethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;$

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;

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biphenyl]-2-sulfonamide;

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N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-
propyl-1-cycloheptimidazolyl)methyl]-2'-(ethoxymethyl)[1,1'-biphenyl]-2-
sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-i)
dimethylisoxazol-5-yl)-2'-(3,3,3-trifluoropropyl) [1,1'-biphenyl]-2-
sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
dimethylisoxazol-5-yl)-2'-(3-fluoropropyl) [1,1'-biphenyl]-2-sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(1,1-
difluoroethyl)-N-(3,4-dimethylisoxazol-5-yl) [1,1'-biphenyl]-2-sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2,2-
trifluoroethyl)-N-(3,4-dimethylisoxazol-5-yl) [1,1'-biphenyl]-2-sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
dimethylisoxazol-5-yl)-2'-[(2-methyl)propoxy] [1,1'-biphenyl]-2-
sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
dimethylisoxazol-5-yl)-2'-(2-methoxyethoxy) [1,1'-biphenyl]-2-sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-
N-(3,4-dimethylisoxazol-5-yl)[1,1'-biphenyl]-2-sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3-
methylisoxazol-5-yl)-2'-trifluoromethyl[1,1'-biphenyl]-2-sulfonamide;
      N-(4-Bromo-3-methyl-5-isoxazolyl)-4'-[(2-butyl-4-oxo-1,3-
diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-trifluoromethyl [1,1'-biphenyl]-2-
sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4-
chloro-3-methyl-5-isoxazolyl)-2'-trifluoromethyl [1,1'-biphenyl]-2-
sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
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dimethyl-5-isoxazolyl)-2'-(N-methoxy-N-methylaminomethyl) [1,1'-

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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2-
            difluoroethoxymethyl)-N-(3,4-dimethylisoxazol-5-yl)[1,1'-biphenyl]-2-
           sulfonamide;
                         4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
            dimethylisoxazol-5-yl)-2'-(2-fluoroethyl)[1,1'-biphenyl]-2-sulfonamide;
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                          4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
            dimethylisoxazol-5-yl)-2'-(2-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;
                          4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
            dimethylisoxazol-5-yl)-2'-(3-methylbutyl)[1,1'-biphenyl]-2-sulfonamide;
                          4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
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            dimethylisoxazol-5-yl)-2'-(2-methypropyl)[1,1'-biphenyl]-2-sulfonamide;
                          4'-[[2-(3,3-Difluorobutyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-
            yl]methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(ethoxymethyl)[1,1'-biphenyl]-
            2-sulfonamide;
                          N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxazolyl]-4'-[[(3-methoxy-2,6-dimethyl-4-isoxa
15
            pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-
            sulfonamide;
                          4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
            dimethylisoxazol-5-yl)-2'-[(1,1-dimethylethoxy)methyl][1,1'-biphenyl]-2-
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            sulfonamide;
                          1-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-
            methoxymethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-
            imidazole-5-carboxamide;
                          1-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-
            methoxymethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-
25
             1H-imidazole-5-carboxamide;
                           1-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-
            biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
                           1-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-
            biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-
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             carboxamide;
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- N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethoxymethyl-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2-fluoroethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;
- $N-(4,5-Dimethyl-3-isoxazolyl)-2'-[(2-fluoroethoxy)methyl]-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] \\ [1,1'-biphenyl]-2-sulfonamide;$
- N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(2-fluoroethoxy)methyl]-4'-[[(3-10 methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2sulfonamide;
 - N-(3,4-Dimethyl-5-isoxazolyl)-2'-ethoxymethyl-4'-[[4-oxo-2-(3,3,3-trifluoropropyl)-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl] [1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1-hydroxyethyl) [1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
 - N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
 - $1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-\\ (ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;$
 - $1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino] sulfonyl-2-\\ (ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;$
- 1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-30 biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

```
methyl][1,1'-biphenyl]-4-yl]methyl]-2-propyl-1H-imidazole-5-carboxamide;
                        1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-
          biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
                        1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-
 5
          biphenyl]-4-yl]methyl]-4-ethyl-N-(1-methylethyl)-2-propyl-1H-imidazole-5-
           carboxamide;
                        1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(2-
           fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-
10
           imidazole-5-carboxamide;
                        N-(4.5-Dimethyl-3-isoxazolyl)-2'-ethoxymethyl-4'-[[(6-ethyl-3-
           methoxy-2-methyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
                        N2-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
           4-yllmethyll-N-methyl-N2-(1-oxobutyl)-L-valinamide;
                        N2-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
15
           4-yl]methyl]-N,N-dimethyl-N2-(1-oxopentyl)-L-valinamide;
                        dimethyl-3-isoxazolyl)-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
                         4'-[(5-Acetyl-4-ethyl-2-propylimidazol-1-yl)methyl]-N-(4,5-dimethyl-
20
           3-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
                        N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethyl-4'-[[(3-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethyl-1-methoxy-2,6-dimethy
           4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
                         N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(6-ethyl-3-methoxy-2-methyl-4-
           pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
25
                         4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-
           N-(4,5-dimethyl-3-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
                         4'-[(2-Buty]-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-4)
           dimethyl-3-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
                         4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-
           N-(4.5-dimethyl-3-isoxazolyl)-2'-fluoro-4-methoxy[1,1'-biphenyl]-2-
30
           sulfonamide;
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N,4-Diethyl-1-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl-2-

${ m HA719f\,CIP}$

	$4'\hbox{-}[(2\hbox{-Butyl-}4\hbox{-}oxo\hbox{-}1,3\hbox{-}diazaspiro[4.4]non-1-en-3-yl)methyl]-3'\hbox{-}chloro-en-3-yl]$
	N-(4,5-dimethyl-3-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
	4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-4)
	dimethyl-3-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
5	$4\text{'-}[(2\text{-Butyl-}4\text{-}oxo\text{-}1,3\text{-}diazaspiro}[4.4] non\text{-}1\text{-}en\text{-}3\text{-}yl) methyl]\text{-}N\text{-}(4,5\text{-}2)$
	dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4-methoxy [1,1'-biphenyl]-2-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4-methoxy [1,1'-biphenyl]-2-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4-methoxy [1,1'-biphenyl]-2-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4-methoxy [1,1'-biphenyl]-2-dimethyl-4-methoxy [1,1'-biphenyl]-4-methoxy [1,1'
	sulfonamide;
	$4\text{'-}[(2\text{-Butyl-}4\text{-}oxo\text{-}1,3\text{-}diazaspiro}[4.4] non\text{-}1\text{-}en\text{-}3\text{-}yl) methyl]\text{-}N\text{-}(4,5\text{-}2)$
	dimethyl-3-is oxazolyl)-3'-fluoro-5-methoxy [1,1'-biphenyl]-2-sulfon a mide;
10	4'-[(2-Butyl-4-oxo-1, 3-diazaspiro[4.4] non-1-en-3-yl) methyl]-5'-chloro-1-2-2-2-2-2-2-2-2-2-2-2-2-2-2-2-2-2-2-
	N-(4,5-dimethyl-3-is oxazolyl)-2'-fluoro-5-methoxy [1,1'-biphenyl]-2-
	sulfonamide;
	$4\text{'-}[(2\text{-Butyl-}4\text{-}oxo\text{-}1,3\text{-}diazaspiro}[4.4]non\text{-}1\text{-}en\text{-}3\text{-}yl)methyl]\text{-}N\text{-}(4,5\text{-}2)$
	dimethyl-3-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
15	4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-4)
	$dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-5-methoxy \cite{1,1'-biphenyl}-2-methoxy 1,1$
	sulfonamide;
	$4\text{'-}[(2\text{-Butyl-}4\text{-}oxo\text{-}1,3\text{-}diazaspiro}[4.4]non\text{-}1\text{-}en\text{-}3\text{-}yl)methyl]\text{-}N\text{-}(4,5\text{-}2)$
	dimethyl-3-isoxazolyl)-3'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-
20	sulfonamide;
	4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-
	N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4,5-dimethoxy [1,1'-biphenyl]-2-dimethyl-3-isoxazolyl)-2'-fluoro-4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4,5-dimethoxy [1,1'-biphenyl]-2-dimethyl-3-isoxazolyl)-2'-fluoro-4,5-dimethoxy [1,1'-biphenyl]-2-dimethyl-3-isoxazolyl)-2'-fluoro-4,5-dimethoxy [1,1'-biphenyl]-2-dimethyl-3-isoxazolyl)-2'-fluoro-4,5-dimethoxy [1,1'-biphenyl]-2-dimethoxy [1,1'-biphenyl]-2-dimethyl-3-dime
	sulfonamide;
	4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-
25	N-(4,5-dimethyl-3-isoxazolyl)-4,5-dimethoxy [1,1'-biphenyl]-2-sulfonamide;
	4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl) methyl]-N-(4,5-4)
	dimethyl-3-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
	$4\text{'-}[(2\text{-Butyl-}4\text{-}oxo\text{-}1,3\text{-}diazaspiro}[4.4]non\text{-}1\text{-}en\text{-}3\text{-}yl)methyl]\text{-}N\text{-}(4,5\text{-}2)$
	dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4, 5-dimethoxy [1,1'-biphenyl]-2-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4, 5-dimethoxy [1,1'-biphenyl]-2-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4, 5-dimethoxy [1,1'-biphenyl]-2-dimethyl-3-dimethoxy [1,1'-biphenyl]-2-dimethyl-3-d
30	sulfonamide;

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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-
            N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;
                           4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-
            N-(4.5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
  5
             dimethyl-3-isoxazolyl)-4-methoxy-3'-methyl [1,1'-biphenyl]-2-sulfonamide;\\
                           4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
             dimethyl-3-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
                           4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
             dimethyl-3-isoxazolyl)-3'-methyl[1,1'-biphenyl]-2-sulfonamide;
10
                           4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
             dimethyl-3-isoxazolyl)-5-fluoro[1,1'-biphenyl]-2-sulfonamide;
                           N<sup>2</sup>-(Cyclopropylcarbonyl)-N<sup>2</sup>-[[2'-[[(4,5-dimethyl-3-
             isoxazolyl)aminolsulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
             valinamide;
15
                           N<sup>2</sup>-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
             vllmethyll-N.3-dimethyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
                           N<sup>2</sup>-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
             vllmethyl]-N-methyl-N<sup>2</sup>-(2-methyl-1-oxopropyl)-L-valinamide;
                           N^2-(Cyclopentylcarbonyl)-N^2-[[2'-[[(4,5-dimethyl-3-
20
             isoxazolyl)amino|sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
             valinamide;
                           N<sup>2</sup>-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl]-3-fluoro[1,1'-
             biphenyll-4-yllmethyll-N-methyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
                           N^2-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
25
             vl]methyl]-N-(1-methylethyl)-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
                           N<sup>2</sup>-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
             yl] methyl]-N-(2-methoxyethyl)-N^2-(1-oxobutyl)-L-valina mide;\\
                           N-(Cyclopropylmethyl)-N<sup>2</sup>-[[2'-[[(4,5-dimethyl-3-
             is oxazolyl) amino] sulfonyl] \\ [1,1'-biphenyl] \\ -4-yl] methyl] -N^2-(1-oxobutyl) -L-vl] \\ [2,1'-biphenyl] \\ -4-yl] \\ [3,1'-biphenyl] \\ -4-yl] \\ [4,1'-biphenyl] \\ -4-yl] \\
30
             valinamide;
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N<sup>2</sup>-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
     yl|methyl|-N<sup>2</sup>-(1-oxobutyl)-N-(3-pyridinyl)-L-valinamide;
            N<sup>2</sup>-[[2'-[[(4.5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
     yllmethyll-N-methyl-N<sup>2</sup>-(1-oxopentyl)-L-valinamide;
            N-Methyl-N<sup>2</sup>-[[2'-[[(5-methyl-3-isoxazolyl)amino]sulfonyl][1,1'-
 5
     biphenyl]-4-yl]methyl]-N<sup>2</sup>-(1-oxopentyl)-L-valinamide;
            N<sup>2</sup>-[[2'-[[(4.5-Dimethyl-3-isoxazolyl)amino|sulfonyl][1,1'-biphenyl]-4-
     yl]methyl]-N-ethyl-N2-(1-oxobutyl)-L-valinamide;
            N<sup>2</sup>-[[2'-[[(4.5-Dimethyl-3-isoxazolyl)amino]sulfonyl]-5'-fluoro[1,1'-
     biphenyl]-4-yl]methyl]-N-methyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
10
            N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(2,6-dimethyl-3-methoxy-4-
      pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
            N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(2,6-dimethyl-3-methoxy-4-
      pyridinyl)oxy|methyl|-3'-fluoro[1,1'-biphenyl]-2-sulfonamide;
15
            N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-
      propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
            N-(4,5-Dimethyl-3-isoxazolyl)-3'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-
      oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
             N-(4,5-Dimethyl-3-isoxazolyl)-5-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-
      oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
20
             4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
      dimethyl-3-isoxazolyl)-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;
             N-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl]-2-
      (ethoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-1-[(1-
      oxopentyl)amino]cyclopentanamide;
25
             and salts, enantiomers and diasteriomer thereof.
```

- 25. A compound of claim 1, wherein said compound is selected from the group consisting of
- 30 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

5

10

20

25

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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(methylamino)methyl][1,1'-biphenyl]-2-sulfonamide;
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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

 $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;$

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3-methyl-2-oxo-1-imidazolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

 $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;$

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

 $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;$

(S)-N-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-[2-methyl-1-(3-methyl-1,2,4-oxadiazol-5-yl)propyl]pentanamide;

N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[2-(2-methoxyethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[2-(ethoxymethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

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4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-
     [(2-oxo-1-pyrrolidinyl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-
     2-sulfonamide;
            4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-
 5
     [(3-methyl-2-oxo-1-imidazolidinyl)methyl]-N-(4,5-dimethyl-3-
     isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
            (S)-2-[N-[2'-[[N-(5-Methyl-3-isoxazolyl)amino]sulfonyl][1,1'-
     biphenyl]-4-yl]methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;
            (S)-2-[N-[2'-[[N-(4-Bromo-5-methyl-3-isoxazolyl)amino]sulfonyl][1,1'-
10
     biphenyl]-4-yl]methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]
     -N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
            4'-[(7-Methoxycarbonyl-2-ethoxybenzimidazol-1-yl)methyl]-
     N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
15
            4'-[(7-Carboxy-2-ethoxybenzimidazol-1-yl)methyl]-N-
     (4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
            4'-[(7-Methoxycarbonyl-2-ethylbenzimidazol-1-yl)methyl]-N-
     (4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
            4'-[(7-Carboxy-2-ethylbenzimidazol-1-yl)methyl]-N-
20
     (4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
            2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-
     (methoxycarbonyl)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-
     isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
            2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-
25
     (carboxy)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-
     biphenyl]-2-sulfonamide;
            2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N-
     methylcarbamoyl)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-
     [1,1'-biphenyl]-2-sulfonamide;
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2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N,N-
     dimethylcarbamoyl)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-
     isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Ethylquinolin-4-yl)oxymethyl]-N-(5-methylisoxazol-3-yl)[1,1'-
 5
     biphenyl]-2-sulfonamide;
            4'-[(5-Acetyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(4,5-
     dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
            4'-[(5-Methoxycarbonyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-
     (4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
10
            4'-[(5-(N,N-dimethylcarbamoyl)-2-n-propyl-4-chloroimidazol-1-
     yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
     dimethyl-3-isoxazolyl)-2'-hydroxymethyl[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
     dimethyl-3-isoxazolyl)-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;
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            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]
     -N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4-
     bromo-5-methyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-
20
     pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
     dimethyl-3-isoxazolyl)-2'-[(formylmethylamino)-methyl][1,1'-biphenyl]-2-
     sulfonamide;
            N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
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     [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-
     methylpropanamide;
            N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
     [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-
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methylcyclopropanecarbox-amide;

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 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,2-dimethylpropanamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2methoxy-N-methylacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethyl-4-pentynamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylcyclobutanecarboxamide;$

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,3-dimethylbutanamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,2,2-trimethylpropanamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3-methoxy-N-methylpropanamide;$

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2-ethoxy-N-methylacetamide;$

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methyl-2-furancarboxamide;$

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 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,4-dimethylpentanamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethyl-3-thiophenecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethylcyclopentaneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethylcyclohexanecarboxamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,3-dimethylbenzamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethylbenzeneacetamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2-fluoro-N-methylbenzamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3fluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-4fluoro-N-methylbenzamide;

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N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-
methylcyclohexaneacetamide;
              N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2-
fluoro-N-methylbenzeneacetamide;
              N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3-
fluoro-N-methylbenzeneacetamide;
              N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-aligned and aligned as a supersymmetric property of the pr
[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-4-
fluoro-N-methylbenzeneacetamide;
              N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-
N,N',N'-trimethylurea;
              N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N'-
(1,1-dimethylethyl)-N-methylurea;
                [[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
\hbox{\tt [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-isoxazolyl)}\\
yl]methyl]methylcarbamic acid ethyl ester;
                \hbox{\tt [[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-}\\
[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
yl]methyl]methylcarbamic acid 2-methylpropyl ester;
              N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-
N,3,3-trimethylbutanamide;
              N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
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[[(4,5-dimethyl-3-isoxazolyl)amino] sulfonyl] [1,1'-biphenyl]-2-yl] methyl]-N-biphenyl] -2-yl] methyl]-N-biphenyl] -2-yl] methyl]-N-biphenyl]-1-yl] methyl]-N-biphenyl]-1-yl] methyl]-N-biphenyl]-1-yl] methyl]-N-biphenyl]-1-yl] methyl]-1-yl] methyl]-N-biphenyl]-1-yl] methyl]-1-yl] methyl]-N-biphenyl]-1-yl] methyl]-1-yl] methyl]-1-yl]-1-yl] methyl]-1-yl

methyl-2-pyridinecarboxamide;

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 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methyl-3-pyridinecarboxamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethyl-2-pyrazinecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,1-dimethyl-1H-pyrrole-2-carboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-Nmethyl-1,2,3-thiadiazole-4-carboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,5-dimethyl-2-pyrazinecarboxamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,3,5-trimethyl-4-isoxazolecarboxamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,3-dimethyl-2-thiophenecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]N,5-dimethyl-2-thiophenecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3cyano-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-4cyano-N-methylbenzamide;

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 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2-methoxy-N-methylbenzamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2-chloro-N-methylbenzamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\ [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3-chloro-N-methylbenzamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-4chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-2,3-difluoro-N-methylbenzamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3,4-difluoro-N-methylbenzamide;$

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-20 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3,5-difluoro-N-methylbenzamide;

4-Acetyl-N-[[4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylbenzamide;

 $N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-\\ [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3-ethoxy-N-methyl-2-thiophenecarboxamide;$

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

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biphenyl]-2-sulfonamide;

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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
 dimethyl-3-isoxazolyl)-2'-[(propylsulfonyl)amino][1,1'-biphenyl]-2-
 sulfonamide;
                              N-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-N-(1-oxopentyl)-L-valine methyl ester;
                              N-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
 yl]methyl]-N-(1-oxopentyl)-L-valine;
                              N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazolyl)-2'-[(3,3-dimethyl-2-isoxazol
 pyrrolidinyl)methyl]-4'-[(2-propyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-
yl)methyl][1,1'-biphenyl]-2-sulfonamide;
                              N-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,
 dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-
 yl]methyl]-N,3,3-trimethylbutanamide;
                              N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-N2-(1-oxopentyl)-L-valinamide;
                              N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
                              4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-2'-[[(2,2,2-trifluoroethyl)amino]methyl][1,1'-
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4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

 $\label{lem:condition} 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid methyl ester;$

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;

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2'-(Cyanomethyl)-N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,7-
     dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-
     sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
     (cyanomethyl)-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
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            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-cyano-
     N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
     dimethyl-3-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
10
            2'-Cyano-N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-
     imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
            N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-methyl[1,1'-
     biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
            N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-
     imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[[(2,2,2-
15
     trifluoroethyl)amino]methyl][1,1'-biphenyl]-2-sulfonamide;
            N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
     [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
     yl]benzeneacetamide;
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            N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
     [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]-3,3-
     dimethylbutanamide;
            2'-Amino-4'-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-
     N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
25
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
     dimethyl-3-isoxazolyl)-2'-nitro[1,1'-biphenyl]-2-sulfonamide;
            N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
     yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-isoleucinamide;
            N2-(Cyclopropylcarbonyl)- N2-[[2'-[[(4,5-dimethyl-3-
     isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
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     isoleucinamide;
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N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-isoleucinamide;
                       N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-isoleucinamide;
                      N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-isoxazolyl) amino[sulfonyl][1,1'-biphenyl]-4-isoxazolyl) amino[sulfonyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphen
yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-isoleucinamide;
                       N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-isoleucinamide;
                       N2\hbox{-}[[2'\hbox{-}[[(4,5\hbox{-}dimethyl\hbox{-}3\hbox{-}isoxazolyl)amino}] sulfonyl][1,1'\hbox{-}biphenyl]\hbox{-}4-
yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-leucinamide;
                       N2-(Cyclopropylcarbonyl)- N2-[[2'-[[(4,5-dimethyl-3-
isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
 leucinamide;
                       N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
 yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-leucinamide;
                        N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-isoxazolyl) amino[sulfonyl][1,1'-biphenyl]-4-isoxazolyl) amino[sulfonyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphenyl][1,1'-biphen
 vl]methyl]-N-methyl- N2-(phenylacetyl)-L-leucinamide;
                        N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
 yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-leucinamide;
                        N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
 yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-leucinamide;
                        N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
 yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-leucinamide;
                        N2-[[2'-[[(4.5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
 yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-valinamide;
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N2-(Cyclopropylcarbonyl)- N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-valinamide;

N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-30 yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-valinamide;

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- N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(phenylacetyl)-L-valinamide;
 N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-valinamide;
 N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
- N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-valinamide;
- N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-valinamide;
- N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-ethyl- N2-(1-oxopentyl)-L-valinamide;
 - N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-hexyl- N2-(1-oxopentyl)-L-valinamide;
 - N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino] sulfonyl]-2-cyano [1,1'-biphenyl]-4-yl] methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-hydroxymethyl[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
 - 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- N,N-Dimethyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
 - N-Methyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
 - N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-30 pyridinyl)oxy]methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

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N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;

2'-Chloro-N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(5,6,7,8-tetrahydro-2-ethyl-4-quinolinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-fluoro[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(phenoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-(Cyclopropylcarbonyl)-N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

N2-(Cyclopropylcarbonyl)- N2-[[2'-[[(4,5-dimethyl-3-

isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

N2-[[2-Chloro-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

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N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2- $(trifluoromethyl) \hbox{$[1,1'$-biphenyl]-4-yl]} methyl]-N-methyl-N2-(1-oxopentyl)-L-methyl-N2-(1-oxopentyl)-N-methyl-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl)-N-methyl-N2-(1-oxopentyl-N2-(1-oxopentyl-N2-(1-oxopentyl-N2-(1-oxopentyl-N2-(1-oxopentyl-N2-(1-oxopentyl-N2-(1-oxopentyl-N2-(1-oxopentyl-N2-(1-oxopentyl-N2-(1-oxo$ valinamide; N2-(Cyclobutylcarbonyl)-N2-[[2'-[[(4,5-dimethyl-3isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-Lvalinamide; 1-[[2-Chloro-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxylic acid; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5dimethyl-3-isoxazolyl)-2'-[(methylsulfonyl)amino][1,1'-biphenyl]-2sulfonamide; (S)-N-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'biphenyl]-4-yl]methyl]-N-[2-methyl-1-[(4-methyl-1piperazinyl)carbonyl]propyl]pentanamide; (S)-N-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'biphenyl]-4-yl]methyl]-N-(1-piperidinyl)carbonyl]propyl]pentanamide; N-(3,3-Dimethylbutyl)- N2-[[2'-[[(4,5-dimethyl-3isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N2-(1-oxopentyl)-Lvalinamide; N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4yl]methyl]-N-[(4-fluorophenyl)methyl]-N2-(1-oxopentyl)-L-valinamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5dimethyl-3-isoxazolyl)-2'-[(methylethoxy)methyl][1,1'-biphenyl]-2sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5dimethyl-3-isoxazolyl)-2'-(propoxymethyl)[1,1'-biphenyl]-2-sulfonamide; 4-Chloro-1-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'biphenyl]-4-yl]methyl]-2-propyl-1H-imidazole-5-carboxamide;

N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(1,2-
     dihydro-2-oxo-1-pyridinyl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-
     biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
     dimethyl-3-isoxazolyl)-2'-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-2-
5
     sulfonamide;
            2-Butyl-4-chloro-1-[[2'-[[(4,5-dimethyl-3-
     isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-1H-imidazole-5-
     carboxamide;
            N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(2-methyl-4-
10
     quinolinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
            N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(2-ethyl-4-quinolinyl)oxy]methyl]
     [1,1'-biphenyl]-2-sulfonamide;
            N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(2-ethyl-5,6,7,8-tetrahydro-4-
     quinolinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
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            N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(2-propyl-4-quinolinyl)oxy]methyl]
     [1,1'-biphenyl]-2-sulfonamide;
            N-(4,5-dimethyl-3-isoxazolyl)-4'-[(5,6,7,8-tetrahydro-2,4-dimethyl-7-
     oxopyrido[2,3-d]pyrimidin-8-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
            4'-[[(2-Ethyl-4-quinolinyl)oxy]methyl]-N-(4,5-dimethyl-3-isoxazolyl)-
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     2'-[(3.3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-
     sulfonamide;
            N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-
     pyrrolidinyl)methyl]-4'-[[(5,6,7,8-tetrahydro-2-ethyl-4-
25
     quinolinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
            3-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
     yl]methyl]-2-ethyl-N-methyl-3H-benzimidazole-4-carboxamide;
            1-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
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yl]methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid phenylmethyl ester; 1-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-phenylethyl ester; 5

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1-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-(2-oxo-1-pyrrolidinyl)ethyl ester;
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1-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 3-(2-oxo-1-pyrrolidinyl)propyl ester;

2'-Cyano-N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(2-ethyl-4-quinolinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

2'-(Cyanomethyl)-N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-10 hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

3-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-5-ethyl-N-methyl-2-propyl-3H-imidazole-4-carboxamide;

1-[[2-Chloro-2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

3-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-5-ethyl-2-propyl-3H-imidazole-4-carboxamide;

3-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethoxy-N-methyl-3H-benzimidazole-4-carboxamide;

3-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethoxy-N,N-dimethyl-3H-benzimidazole-4-carboxamide;

2-[[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]propylamino]-3-pyridinecarboxylic acid;

4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

 $4\text{'-}[(2,7\text{-}Diethyl\text{-}5H\text{-}pyrazolo[1,5\text{-}b][1,2,4]triazol\text{-}5\text{-}yl)methyl]\text{-}N\text{-}(4,5\text{-}dimethyl\text{-}3\text{-}isoxazolyl)[1,1\text{'-}biphenyl]\text{-}2\text{-}sulfonamide;}$

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sulfonamide:

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N-[2-Butyl-3-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-
biphenyl]-4-yl]methyl]-3,4-dihydro-4-oxo-6-quinazolinyl]-N'-methyl-N'-(1-
methylethyl)urea;
      2-[[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]propylamino]-N-methyl-3-pyridinecarboxamide;
      4'-[[(3-Methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-N-(4,5-
dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
      N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-
pyrrolidinyl)methyl]-4'-[[(3-methoxy-2,6-dimethyl-4-
pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
      N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-
pyrrolidinyl)methyl]-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-
cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
      N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-
pyridinyl)oxy]methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
      3-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
yl]methyl]-2-ethyl-5-methyl-3H-imidazole-4-carboxamide;
      4'-[(2-Butyl-3,4-dihydro-4-oxo-3-quinazolinyl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-
biphenyl]-2-sulfonamide;
      N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-[(3,3-
dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-4-yl]methyl]-N-
methyl-N2-(1-oxopentyl)-L-valinamide;
      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-2'-[(4,4-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-
biphenyl]-2-sulfonamide;
      4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(4,5-dimethyl-3-
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isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-

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- N-[2-[[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl]methyl]methylamino]ethyl]acetamide;
- 4-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-5 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-acetic acid ethyl ester;
 - N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N2-(1-oxopentyl)-N-propyl-L-valinamide;
- N2-[[2'-[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-410 yl]methyl]-N2-(1-oxopentyl)-N-[(tetrahydro-2-furanyl)methyl]-Lvalinamide;
 - 2'-Chloro-N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(2-ethyl-4-quinolinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
 - N-(4,5-dimethyl-3-isoxazolyl)-4'-[[(2-ethyl-4-quinolinyl)oxy]methyl]-2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-chloro-N-(4.5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-methylpropoxy)methyl][1,1'-biphenyl]-2-sulfonamide;
 - $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(ethylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;$
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-25 dimethyl-3-isoxazolyl)-2'-[(2,2,2-trifluoroethoxy)methyl][1,1'-biphenyl]-2sulfonamide;
 - $\label{lem:conditional} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-fluoroethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;$
- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;

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N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-
propyl-1-cycloheptimidazolyl)methyl]-2'-(ethoxymethyl)[1,1'-biphenyl]-2-
sulfonamide;
                      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-4)
dimethyl-3-isoxazolyl)-2'-(3,3,3-trifluoropropyl)\ [1,1'-biphenyl]-2-isoxazolyl-2'-(3,3,3-trifluoropropyl)\ [1,1'-biphenyl]-2'-(3,3,3-trifluoropropyl)\ [1,1'-b
sulfonamide;
                      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-in-1)methyl]
dimethylisoxazol-3-yl)-2'-(3-fluoropropyl)\ [1,1'-biphenyl]-2-sulfonamide;
                      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(1,1-
difluoroethyl)-N-(4,5-dimethylisoxazol-3-yl) [1,1'-biphenyl]-2-sulfonamide;
                      4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2,2-
trifluoroethyl)-N-(4,5-dimethylisoxazol-3-yl) [1,1'-biphenyl]-2-sulfonamide;
                       4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
 dimethylisoxazol-3-yl)-2'-[(2-methyl)propoxy] [1,1'-biphenyl]-2-
 sulfonamide;
                       4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
 dimethylisoxazol-3-yl)-2'-(2-methoxyethoxy) [1,1'-biphenyl]-2-sulfonamide;
                       4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl-2'-butyl-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1'-[(
 N-(4,5-dimethylisoxazol-3-yl)[1,1'-biphenyl]-2-sulfonamide;
                       4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
 dimethyl-3-isoxazolyl)-2'-(N-methoxy-N-methylaminomethyl) [1,1'-
 biphenyl]-2-sulfonamide;
                       4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2-
  difluoroethoxymethyl)-N-(4,5-dimethylisoxazol-3-yl)[1,1'-biphenyl]-2-
 sulfonamide;
                       4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-4)
  dimethylisoxazol-3-yl)-2'-(2-fluoroethyl)[1,1'-biphenyl]-2-sulfonamide;
                        4'-[(2-Buty]-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-y])methyl]-N-(4,5-4)
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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(3-methylbutyl)[1,1'-biphenyl]-2-sulfonamide;

dimethylisoxazol-3-yl)-2'-(2-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

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 $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(2-methypropyl)[1,1'-biphenyl]-2-sulfonamide;$

 $\label{lem:condition} 4'-[[2-(3,3-Difluorobutyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;$

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-[(1,1-dimethylethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

N,4-Diethyl-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino] sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-(1-methylethyl)-2-propyl-1H-imidazole-5-carboxamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-ethoxymethyl-4'-[[(6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;

4'-[(5-Acetyl-4-ethyl-2-propylimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(6-ethyl-3-methoxy-2-methyl-4pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide; 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4dimethyl-5-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro-4-methoxy[1,1'-biphenyl]-2sulfonamide; 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4dimethyl-5-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-15 dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)-4-methoxy[1,1'-biphenyl]-2sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4dimethyl-5-isoxazolyl)-3'-fluoro-5-methoxy[1,1'-biphenyl]-2-sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-20 N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro-5-methoxy[1,1'-biphenyl]-2sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4dimethyl-5-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-25 dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)-5-methoxy[1,1'-biphenyl]-2sulfonamide; 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4dimethyl-5-isoxazolyl)-3'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2sulfonamide;

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4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-
      N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-
      sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-
 5
      N-(3,4-dimethyl-5-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
      dimethyl-5-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
      dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)-4,5-dimethoxy[1,1'-biphenyl]-2-
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      sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-
      N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-
      N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
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            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
      dimethyl-5-isoxazolyl)-4-methoxy-3'-methyl[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
      dimethyl-5-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
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      dimethyl-5-isoxazolyl)-3'-methyl[1,1'-biphenyl]-2-sulfonamide;
            4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
      dimethyl-5-isoxazolyl)-5-fluoro[1,1'-biphenyl]-2-sulfonamide;
            N<sup>2</sup>-(Cyclopropylcarbonyl)-N<sup>2</sup>-[[2'-[[(3,4-dimethyl-5-
      isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
25
      valinamide;
            N^2-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
      yl]methyl]-N,3-dimethyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
            N<sup>2</sup>-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
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yl]methyl]-N-methyl-N²-(2-methyl-1-oxopropyl)-L-valinamide;

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N<sup>2</sup>-(Cyclopentylcarbonyl)-N<sup>2</sup>-[[2'-[[(3,4-dimethyl-5-
      isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
      valinamide;
             N<sup>2</sup>-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-3-fluoro[1,1'-
      biphenyl]-4-yl]methyl]-N-methyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
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             N<sup>2</sup>-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
      vllmethyll-N-(1-methylethyl)-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
             N<sup>2</sup>-[[2'-[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
      yllmethyll-N-(2-methoxyethyl)-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
             N-(Cyclopropylmethyl)-N<sup>2</sup>-[[2'-[[(3,4-dimethyl-5-
10
      isoxazolyl)amino|sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N<sup>2</sup>-(1-oxobutyl)-L-
      valinamide;
             N<sup>2</sup>-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
      yl]methyl]-N<sup>2</sup>-(1-oxobutyl)-N-(3-pyridinyl)-L-valinamide;
             N<sup>2</sup>-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
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      yllmethyll-N-methyl-N<sup>2</sup>-(1-oxopentyl)-L-valinamide;
             N-Methyl-N<sup>2</sup>-[[2'-[[(3-methyl-5-isoxazolyl)amino]sulfonyl][1,1'-
      biphenyl]-4-yl]methyl]-N<sup>2</sup>-(1-oxopentyl)-L-valinamide;
             N<sup>2</sup>-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
      yl]methyl]-N-ethyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
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             N<sup>2</sup>-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-5'-fluoro[1,1'-
      biphenyl]-4-yl]methyl]-N-methyl-N<sup>2</sup>-(1-oxobutyl)-L-valinamide;
             N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2,6-dimethyl-3-methoxy-4-
      pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
             N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2,6-dimethyl-3-methoxy-4-
25
      pyridinyl)oxy]methyl]-3'-fluoro[1,1'-biphenyl]-2-sulfonamide;
             N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-
      propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
             N-(3,4-Dimethyl-5-isoxazolyl)-3'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-
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      oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
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N-(3,4-Dimethyl-5-isoxazolyl)-5-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;
N-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(ethoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-1-[(1-oxopentyl)amino]cyclopentanamide;
and salts, enantiomers and diasteriomer thereof.

- 26. A method for the treatment of an endothelin-dependent or angiotensin II-dependent disorder, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
- 27. A method for the treatment of hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
- 28. A method for the treatment of pulmonary hypertension,comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
 - 29. A method for the treatment of primary pulmonary hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
 - 30. A method for the treatment of low renin hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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- 31. A method for the treatment of male erectile dysfunction, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
- 5 32. A method for the treatment of male or female sexual dysfunction, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
- 33. A method for the treatment of heart failure, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
 - 34. A method for the treatment of atherosclerosis, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
 - 35. A method for the treatment of restenosis, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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36. A method for the treatment of endotoxemia, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

- 37. A method for the inhibition of cell growth, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
- 38. A method for the treatment of cancer, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

39. A method for the treatment of migraine, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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40. A method for the treatment of asthma, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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41. A method for the treatment of ischemia, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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42. A method for the treatment of subarachnoid hemorrhage, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

43. A method for the treatment of benign prostatic hypertrophy, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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44. A method for the treatment of renal, glomerular or mesangial cell disorders, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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45. A method for the treatment of acute or chronic renal failure, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

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46. A pharmaceutical composition for the treatment of an endothelin-dependent or angiotensin II-dependent disorder, comprising a

pharmaceutically acceptable vehicle or diluent and at least one compound of claim 1 in an amount effective therefor.

47. A compound of the formula

5 wherein

 $R_{\scriptscriptstyle 2}$, $R_{\scriptscriptstyle 3}$, $R_{\scriptscriptstyle 101}$, $R_{\scriptscriptstyle 102}$, $R_{\scriptscriptstyle 103}$, and $R_{\scriptscriptstyle 104}$ are as defined in claim 1; $R_{\scriptscriptstyle 50}$ is hydroxy, chloro, bromo, iodo, -OSO $_{\scriptscriptstyle 2}$ -alkyl, or -OSO $_{\scriptscriptstyle 2}$ -aryl; and $R_{\scriptscriptstyle 51}$ is hydrogen, or a suitable nitrogen protecting group.

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- 48. The compound of claim 47, wherein R_{51} is $-CH_2OCH_2CH_2OCH_3, -CH_2OCH_2CH_2Si(CH_3)_3, -CH_2OCH_3, \text{ or } -CH_2OCH_2-\text{aryl}.$
 - 49. A compound of the formula

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wherein R_1 , R_2 , R_3 , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1; and R_{51} is a suitable nitrogen protecting group.

- 50. The compound of claim 49, wherein R₅₁ is -CH₂OCH₂CH₂OCH₃, -CH₂OCH₂CH₂Si(CH₃)₃, -CH₂OCH₃, or -CH₂OCH₂-aryl.
 - 51. A compound of the formula

$$\begin{array}{c|c} R_1 \\ R_{101} \\ R_2 \\ R_{52} \end{array} \qquad \text{(LVII)}$$

wherein

 $\boldsymbol{R}_{_{1}}\!,\,\boldsymbol{R}_{_{2}\!,}\boldsymbol{R}_{_{101}}$ and $\boldsymbol{R}_{_{102}}$ are as defined in claim 1; and

 $5 \qquad R_{\rm 52} \ is \ chloro, \ bromo, \ iodo, \ or \ -OSO_2CF_3.$

52. A compound of claim 51

wherein

- 53. A process for the preparation of a compound of claim 1, comprising at least one of the following steps:
- 15 a) displacement of a leaving group R_{50} via the anion of a compound $R_1\text{-H}$ from a compound of the formula

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wherein R₁, R₂, R₃, R₁₀₁, R₁₀₂, R₁₀₃, and R₁₀₄ are as defined in claim 1, R₅₀ is hydroxy, chloro, bromo, iodo, -OSO₂-alkyl or -OSO₂-aryl, and R₅₁ is hydrogen or a suitable nitrogen protecting group, using a Mitsunobu reaction or S_n1 or S_n2 displacement reaction, with removal of said nitrogen protecting group as appropriate;

b) removal of the nitrogen protecting group $R_{\scriptscriptstyle{51}}$ from a compound of formula

- wherein R_1 , R_2 , R_{3} , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1, and R_{51} is a suitable nitrogen protecting group;
 - c) organometallic coupling of a compound of formula

with a compound of formula

$$R_{103}$$
 R_{104}
 R_{104}
 R_{104}
 R_{104}
 R_{104}
 R_{104}
 R_{104}
 R_{104}
 R_{104}

wherein

- $R_{_{1}}$, $R_{_{2}}$, $R_{_{3,}}$, $R_{_{101}}$, $R_{_{102}}$, $R_{_{103}}$, and $R_{_{104}}$ are as defined for a compound of formula I and $R_{_{51}}$ is a suitable nitrogen protecting group;
- R_{52} is chloro, bromo, iodo or $-OSO_2CF_3$ and R_{54} is a boronic acid, boronic ester or stannane derivative;

or R_{52} is a boronic acid, boronic ester or stannane derivative and R_{54} is chloro, bromo, iodo or $-OSO_2CF_3$;

d) acylation of a compound of the formula

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wherein R_1 , R_{3} , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1, and R_{51} is hydrogen or a suitable nitrogen protecting group, with an acylating agent of the formula R_{55} -(C=O) R_{22} , R_{19} N=C=O, R_{55} -CO $_2$ R $_{18}$, R_{55} SO $_2$ R $_{22}$, wherein R_{18} , R_{19} and R_{22} are as defined in claim 1 and R_{55} is an activating group for an acid, or made using an acid activating agent, with removal of said nitrogen protecting group as appropriate; or

e) reductive amination of a compound of formula

wherein R_1 , R_3 , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1, and R_{51} is hydrogen or a suitable nitrogen protecting group, with an amine of the formula

wherein R_{23} , R_{24} , and x are as defined in claim 1, with removal of said nitrogen protecting group as appropriate.

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54. A compound of claim 1, wherein said compound is selected from the group consisting of:

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-ethyl[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(2,2-dimethylpropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(2-ethoxyethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-ethyl[1,1'-biphenyl]-2-sulfonamide;

 $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2,2-dimethylpropyl)[1,1'-biphenyl]-2-sulfonamide;$

 $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2-ethoxyethyl)[1,1'-biphenyl]-2-sulfonamide;$

N-(3,4-Dimethyl-5-isoxazolyl)-2'-ethyl-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

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N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
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2'-[(1,1-Dimethylethoxy)methyl]- N-(3,4-dimethyl-5-isoxazolyl)-4'[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethyl-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]- N-(3,4-dimethyl-5-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2sulfonamide;

 $N-(4,5-Dimethyl-4-isoxazolyl)-2'-ethyl-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl) methyl] \\ [1,1'-biphenyl]-2-sulfonamide;$

N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-

[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-4'[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'biphenyl]-2-sulfonamide;

 $N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethyl-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide; \\N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-$

30 [(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

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- N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 2'-[(1,1-Dimethylethoxy)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-4'[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'biphenyl]-2-sulfonamide
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-10 dimethyl-5-isoxazolyl)-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;
 - 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;
 - N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;
 - N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;
 - $N-(3,4-Dimethyl-5-isoxazolyl)-\ 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;$
 - N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-30 propyl-1-cycloheptimidazolyl)methyl]-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;

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- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;
 - $\label{lem:condition} 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;$
- N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;
 - N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;
 - $N-(4,5-Dimethyl-3-isoxazolyl)-\ 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;$
- N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;
 - N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;
 - N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-30 pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2sulfonamide;

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- $N-(4,5-Dimethyl-3-isoxazolyl)-\ 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;$
- N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;
 - N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
 - N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
 - N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
 - N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
 - N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
 - $N-(3,4-Dimethyl-5-isoxazolyl)-\ 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;$
- 4'-[[2-(2,2-Difluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-30 yl]methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

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4'-[[2-(2,2-Difluor obutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

 $\label{lem:conditional} 4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;$

4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

 $\label{lem:condition} 4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;$

4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

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N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;
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N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(2-fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

 $1\hbox{-}[[2'\hbox{-}[[(3,4\hbox{-}Dimethyl\hbox{-}5\hbox{-}isoxazolyl)amino]sulfonyl\hbox{-}2\hbox{-}$

25 (ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino] sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl] methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

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biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-
     carboxamide;
            1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(2-
 5
     fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-
     propyl-1H-imidazole-5-carboxamide;
            1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-
     (ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-
     imidazole-5-carboxamide;
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            1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-
     biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
            1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-
     biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
            1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(2-
15
     fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-
     imidazole-5-carboxamide;
            1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-
     (ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-
     5-carboxamide;
20
            N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(6-ethyl-3-methoxy-2-methyl-4-
     pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
            N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(6-ethyl-3-methoxy-2-methyl-4-
     pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
           N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2-methyl-6-propyl-4-
25
     pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
           N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-methoxy-2-methyl-6-propyl-4-
      pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
           N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2-methyl-6-propyl-4-
     pyridinyl)oxy]methyl]-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;
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           N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-methoxy-2-methyl-6-propyl-4-
     pyridinyl)oxy]methyl]-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;
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1-[[2'-[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-

	N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2-methyl-6-propyl-4-methyl-6-propyl-
	pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
	N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-methoxy-2-methyl-6-propyl-4-methyl-6-propyl-
	pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
5	N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-ethoxy-6-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3
	pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
	N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-ethoxy-6-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazoly
	pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
	N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-ethoxy-6-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3-ethyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3
10	pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
	N-(3,4-Dimethyl-5-is oxazolyl)-4'-[[(3-ethoxy-6-ethyl-2-methyl-4-is oxazolyl]-4'-[[(3-ethoxy-6-ethyl-2-methyl-4-is oxazolyl]-4'-[[(3-ethoxy-6-ethyl-4-is oxazolyl]-4'-[[(3-ethoxy-6-ethyl-4
	pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
	N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-ethoxy-6-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethoxy-6-ethyl-4-isoxazolyl)-4'-[(3
	pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
1 5	N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-ethoxy-6-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-2-methyl-4-isoxazolyl)-4'-[(3-ethyl-4-
	pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
	$N\hbox{-}(4,5\hbox{-}Dimethyl\hbox{-}3\hbox{-}isoxazolyl)\hbox{-}4'\hbox{-}[[(3\hbox{-}ethoxy\hbox{-}2\hbox{-}methyl\hbox{-}6\hbox{-}propyl\hbox{-}4\hbox{-}$
	pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
	N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-ethoxy-2-methyl-6-propyl-4-methyl-6-propyl-6
20	pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
	$N\hbox{-}(4,5\hbox{-}Dimethyl\hbox{-}3\hbox{-}isoxazolyl)\hbox{-}4'\hbox{-}[[(3\hbox{-}ethoxy\hbox{-}2\hbox{-}methyl\hbox{-}6\hbox{-}propyl\hbox{-}4\hbox{-}4]]}$
	pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
	$N\hbox{-}(3,4\hbox{-}Dimethyl\hbox{-}5\hbox{-}isoxazolyl)\hbox{-}4'\hbox{-}[[(3\hbox{-}ethoxy\hbox{-}2\hbox{-}methyl\hbox{-}6\hbox{-}propyl\hbox{-}4\hbox{-}2]]}$
	pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
25	N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-ethoxy-2-methyl-6-propyl-4-methyl-6-propyl-6
	pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
	N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(3-ethoxy-2-methyl-6-propyl-4-methyl-6-propyl-6
	pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
	$N\hbox{-}(4,5\hbox{-}Dimethyl\hbox{-}3\hbox{-}isoxazolyl)\hbox{-}4'\hbox{-}[[(2,6\hbox{-}dimethyl\hbox{-}3\hbox{-}ethoxy\hbox{-}4\hbox{-}4]$
30	pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2,6-dimethyl-3-ethoxy-4-

pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide; N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(2,6-dimethyl-3-ethoxy-4-

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide; and salts, enantiomers and diasteriomers thereof.

55. A compound of formula LXX, or an enantiomer, diastereomer or salt thereof

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wherein R_2 , R_{101} , R_{102} , R_{103} , R_{104} are as defined in claim 1;

 ${
m R_3}$ is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen; and

 $R_{_{1}}$ is any group such that the resulting compound demonstrates affinity $(IC_{_{50}}) \ for \ both \ the \ AT_{_{1}} \ receptor \ and \ ET_{_{A}} \ receptor \ of \ less \ than \ 5$ micromolar at both receptors.

56. A compound of claim 5, wherein R₁ is

salt, enantiomer or diasteriomer thereof.

- 66. N-(4,5-Dimethyl-3-isoxazolyl)-2'-[(2-fluoroethoxy)methyl]-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diasteriomer thereof.
- 67. N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diasteriomer thereof.

$$\stackrel{\mathsf{R}_6}{\stackrel{\mathsf{N}}{\longrightarrow}} \stackrel{\mathsf{R}_9}{\stackrel{\mathsf{R}_7}{\longrightarrow}}$$

68. A compound of claim 5, where in R₁ is

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- 69. A compound of claim 68, wherein R_2 is hydrogen, alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R_{101} , R_{102} , R_{103} , R_{104} are each independently hydrogen, halogen, or alkyl.
- 15 70. A compound of claim 68 wherein R_2 is $-CH_2Y$.
 - 71. A compound of claim 70, wherein Y is Q.
- 72. A method for the treatment of chronic obstructive pulmonary disease, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
 - 73. A method for the alleviation of pain associated with prostate cancer, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
 - 74. A method for preventing or reducing organ damage associated with the cell poliferative effects of endothelin, comprising the step of

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administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

- 75. A method for reducing general morbidity and mortality associated with endothelin-dependent or angiotensin II-dependent disorders, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
- 76. A method for the treatment of diabetic nephropathy, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
 - 77. A method for the treatment of dementia, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.
 - 78. A compound of claim 1 wherein R_3 is other than pyridyl when R_1 is \underline{A} .
- 79. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one ACE inhibitor.
- 80. The method of claim 79 wherein said ACE inhibitor is selected from captopril, zofenopril, fosinopril, ceranapril, alacepril, enalapril, delapril, pentopril, quinapril, ramipril, or lisinopril.
 - 81. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one vasopepsidase inhibtor.

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- 82. The method of claim 81 wherein said vasopepsidase inhibitor is selected from omapatrilat or gemopatrilat.
- 83. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one HMG CoA reductase inhibitor.
 - 84. The method of claim 83 wherein said HMG CoA reductase inhibitor is selected from pravastatin, lovastatin, atorvastatin, simvastatin, NK-104 or ZD-4522.
 - 85. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one anti-platelet agent.
 - 86. The method of claim 85 wherein said anti-platelet agent is selected from clopidigrel, ticlopidine, CS-747 or aspirin.
 - 87. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one anti-diabetic agent.
 - 88. The method of claim 87 wherein said anti-diabetic agent is selected from biguanides or biguanide/glyburide combinations.
 - 89. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one beta-adrenergic agent.

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- 90. The method of claim 89 wherein said beta-adrenergic agent is selected from carvedilol or metoprolol.
- 91. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one mineralocorticoid receptor antagonist.
 - 92. The method of claim 91 wherein said mineralocorticoid receptor antagonist is selected from spironolactone or eplerenone.
 - 93. The pharmaceutical composition of claim 46 further comprising at least one ACE inhibitor.
- 94. The pharmaceutical composition of claim 93 wherein said
 15 ACE inhibitor is selected from captopril, zofenopril, fosinopril,
 ceranapril, alacepril, enalapril, delapril, pentopril, quinapril, ramipril,
 or lisinopril.
- 95. The pharmaceutical composition of claim 46 further comprising at least one vasopepsidase inhibitor.
 - 96. The pharmaceutical composition of claim 95 wherein said vasopepsidase inhibitor is selected from omapatrilat or gemopatrilat.
- 97. The pharmaceutical composition of claim 46 further comprising at least one HMG CoA reductase inhibitor.
 - 98. The pharmaceutical composition of claim 97 wherein said HMG CoA reductase inhibitor is selected from pravastatin, lovastatin, atorvastatin, simvastatin, NK-104 or ZD-4522.

- 99. The pharmaceutical composition of claim 46 further comprising at least one anti-platelet agent.
- 100. The pharmaceutical composition of claim 99 wherein said anti-platelet agent is selected from clopidigrel, ticlopidine, CS-747 or aspirin.
- 101. The pharmaceutical composition of claim 46 further10 comprising at least one anti-diabetic agent.
 - 102. The pharmaceutical composition of claim 101 wherein said anti-diabetic agent is selected from biguanides or biguanide/glyburide combinations.

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- 103. The pharmaceutical composition of claim 46 further comprising at least one beta-adrenergic agent.
- 104. The pharmaceutical composition of claim 103 wherein said beta-adrenergic agent is selected from carvedilol or metoprolol.
 - 105. The pharmaceutical composition of claim 46 further comprising at least one mineralocorticoid receptor antagonist.
- 25 106. The pharmaceutical composition of claim 105 wherein said mineralocorticoid receptor antagonist is selected from spironolactone or eplerenone.

- 107. The compound of claim 62 that is (+)-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'- (ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide.
- 5 108. The compound of claim 62 that is (-)-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'- (ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide.